Time stamp	2004/03/12 12:05	2004/03/12 12:06	2004/03/12 12:06	2004/03/12 12:07	2004/03/12 12:10	2004/03/12 12:07		2004/03/12 12:07			2004/03/12 12:10
DB	USPAT	USPAT	USPAT	USPAT	USPAT	USPAT	4.	USPAT			USPAT
its Search Text	0 ('514/183,,243,63,408,416,417").CCLS		[287] ("514/408,416,417").ccLs	704 ("548/452,470,472").ccls	86 ("514/183,13,243").ccLS) and ("514/408,416,417").ccLS)	("548/452,470,472").CCLS) and (("514/183	(("514/408,416,417").ccls))	9 ("514/183,13,243").CCLS) and (("514/408,416,417").CCLS) and	("548/452,470,472").CCLS) and (("514/183,13,243").CCLS) and	(("514/408,416,417").ccls))	11 (("514/183,13,243").CCLS) and (("514/408,416,417").CCLS) and respiratory
Hits		3709	128	70.	α	•					7
I Number		2	m	4	·	9		7			80

```
Welcome to STN International! Enter x:x
LOGINID:ssspta1611sxp
```

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

```
Welcome to STN International
NEWS 1
                 Web Page URLs for STN Seminar Schedule - N. America
NEWS 2
                 "Ask CAS" for self-help around the clock
NEWS \, 3 SEP 09 CA/CAplus records now contain indexing from 1907 to the
                present
NEWS 4 DEC 08 INPADOC: Legal Status data reloaded
NEWS 5 SEP 29 DISSABS now available on STN
NEWS 6 OCT 10 PCTFULL: Two new display fields added
NEWS 7 OCT 21 BIOSIS file reloaded and enhanced
NEWS 8 OCT 28 BIOSIS file segment of TOXCENTER reloaded and enhanced
NEWS 9 NOV 24 MSDS-CCOHS file reloaded
NEWS 10 DEC 08 CABA reloaded with left truncation
NEWS 11 DEC 08 IMS file names changed
NEWS 12 DEC 09 Experimental property data collected by CAS now available
                 in REGISTRY
NEWS 13 DEC 09 STN Entry Date available for display in REGISTRY and CA/CAplus
NEWS 14 DEC 17 DGENE: Two new display fields added
NEWS 15 DEC 18 BIOTECHNO no longer updated
NEWS 16 DEC 19
                CROPU no longer updated; subscriber discount no longer
                available
NEWS 17 DEC 22 Additional INPI reactions and pre-1907 documents added to CAS
                databases
NEWS 18 DEC 22 IFIPAT/IFIUDB/IFICDB reloaded with new data and search fields
NEWS 19 DEC 22 ABI-INFORM now available on STN
NEWS 20 JAN 27 Source of Registration (SR) information in REGISTRY updated
                and searchable
NEWS 21 JAN 27 A new search aid, the Company Name Thesaurus, available in
                CA/CAplus
NEWS 22 FEB 05
                German (DE) application and patent publication number format
                changes
NEWS 23 MAR 03 MEDLINE and LMEDLINE reloaded
NEWS 24 MAR 03 MEDLINE file segment of TOXCENTER reloaded
NEWS 25 MAR 03 FRANCEPAT now available on STN
NEWS EXPRESS MARCH 5 CURRENT WINDOWS VERSION IS V7.00A, CURRENT
             MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP),
             AND CURRENT DISCOVER FILE IS DATED 3 MARCH 2004
NEWS HOURS
             STN Operating Hours Plus Help Desk Availability
NEWS INTER
             General Internet Information
NEWS LOGIN
             Welcome Banner and News Items
NEWS PHONE
             Direct Dial and Telecommunication Network Access to STN
NEWS WWW
             CAS World Wide Web Site (general information)
```

Enter NEWS followed by the item number or name to see news on that specific topic.

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FILE 'HOME' ENTERED AT 10:33:14 ON 12 MAR 2004

=> file reg
COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 10:33:47 ON 12 MAR 2004 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2004 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 10 MAR 2004 HIGHEST RN 661450-61-9 DICTIONARY FILE UPDATES: 10 MAR 2004 HIGHEST RN 661450-61-9

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 6, 2004

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

=>
Uploading c:\program files\stnexp\queries\09869668.1

L1 STRUCTURE UPLOADED

=> d 11

L1 HAS NO ANSWERS

L1 STR

Patel

<3/12/2004>

G1 O, OH, MeO, EtO, n-PrO, i-PrO, n-BuO, i-BuO, s-BuO, t-BuO, X

G2 CF3, C (O) CH3

G3 CH,Cb

Structure attributes must be viewed using STN Express query preparation.

=> s 11

SAMPLE SEARCH INITIATED 10:34:19 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED - 17055 TO ITERATE

5.9% PROCESSED 1000 ITERATIONS INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED) SEARCH TIME: 00.00.01

0 ANSWERS

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS:

333286 TO 348914

PROJECTED ANSWERS:

OT 0

0 SEA SSS SAM L1

=> s ll sss full

FULL SEARCH INITIATED 10:34:30 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED - 343745 TO ITERATE

100.0% PROCESSED 343745 ITERATIONS SEARCH TIME: 00.00.11

2 ANSWERS

L3

2 SEA SSS FUL L1

=> file marpat

COST IN U.S. DOLLARS

SINCE FILE

ENTRY

TOTAL SESSION 156.05

FULL ESTIMATED COST

155.84

FILE 'MARPAT' ENTERED AT 10:34:49 ON 12 MAR 2004 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

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FILE CONTENT: 1988-PRESENT (VOL 140 ISS 10) (20040307/ED)

MOST RECENT CITATIONS FOR PATENTS FROM FIVE MAJOR ISSUING AGENCIES (COVERAGE TO THESE DATES IS NOT COMPLETE):

US 6683216 27 JAN 2004 DE 10317487 12 FEB 2004 EP 1388563 11 FEB 2004 JP 2004047131 12 FEB 2004 WO 2004011964 05 FEB 2004

Structure search limits have been raised. See HELP SLIMIT for the new, higher limits.

=> s 11 sss full FULL SEARCH INITIATED 10:35:18 FILE 'MARPAT' FULL SCREEN SEARCH COMPLETED - 70092 TO ITERATE

8.3	3 ક	PROCESSED	5836	ITERATIONS	(1	INCOMPLETE)	1	ANSWERS
11.9	8	PROCESSED	8339	ITERATIONS	(3	INCOMPLETE)	3	ANSWERS
21.7	7 용	PROCESSED	15225	ITERATIONS	(6	INCOMPLETE)	8	ANSWERS
28.0) &	PROCESSED	19632	ITERATIONS	(14	INCOMPLETE)	. 16	ANSWERS
33.8	3 %	PROCESSED	23702	ITERATIONS	(22	INCOMPLETE)	24	ANSWERS
40.8	3 €	PROCESSED	28566	ITERATIONS	(36	INCOMPLETE)	39	ANSWERS
47.1	1 ક	PROCESSED	33045	ITERATIONS	(44	INCOMPLETE)	47	ANSWERS
54.2	28	PROCESSED	37970	ITERATIONS	(55	INCOMPLETE)	58	ANSWERS
59.4	8	PROCESSED	41651	ITERATIONS	(62	INCOMPLETE)	65	ANSWERS
65.4	1 %	PROCESSED	45833	ITERATIONS	(73	INCOMPLETE)	76	ANSWERS
70.4	1 ક	PROCESSED	49356	ITERATIONS	(81	INCOMPLETE)	84	ANSWERS
75.2	28	PROCESSED	52680	ITERATIONS	(95	INCOMPLETE)	98	ANSWERS
79.1	L &	PROCESSED	55444	ITERATIONS	(106	INCOMPLETE)	109	ANSWERS
81.5	કે	PROCESSED	57097	ITERATIONS	(111	INCOMPLETE)	114	ANSWERS
82.6	5 ક	PROCESSED	57914	ITERATIONS	(115	INCOMPLETE)	118	ANSWERS
84.1	ક	PROCESSED	58954	ITERATIONS	(122	INCOMPLETE)	125	ANSWERS
85.0) 웅	PROCESSED	59580	ITERATIONS	(122	INCOMPLETE)	125	ANSWERS
85.4	ક	PROCESSED	59879	ITERATIONS	(123	INCOMPLETE)	126	ANSWERS
85.6	58	PROCESSED	60000	ITERATIONS	(123	INCOMPLETE)	126	ANSWERS

Patel <3/12/2004>

INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED) SEARCH TIME: 00.05.13

FULL FILE PROJECTIONS: ONLINE **INCOMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS:

70092 TO 70092

PROJECTED ANSWERS:

126 TO 183

L4 126 S

126 SEA SSS FUL L1

=> file caplus

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 113.20 269.25

FULL ESTIMATED COST

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FILE COVERS 1907 - 12 Mar 2004 VOL 140 ISS 12 FILE LAST UPDATED: 11 Mar 2004 (20040311/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d his

(FILE 'HOME' ENTERED AT 10:33:14 ON 12 MAR 2004)

FILE 'REGISTRY' ENTERED AT 10:33:47 ON 12 MAR 2004

L1 STRUCTURE UPLOADED

L2 0 S L1

L3 2 S L1 SSS FULL

FILE 'MARPAT' ENTERED AT 10:34:49 ON 12 MAR 2004 L4 126 S L1 SSS FULL

FILE 'CAPLUS' ENTERED AT 10:40:39 ON 12 MAR 2004

=> d 13 fbib hitstr abs total
YOU HAVE REQUESTED DATA FROM FILE 'REGISTRY' - CONTINUE? (Y)/N:y

'FBIB' IS NOT A VALID FORMAT FOR FILE 'REGISTRY'

'HITSTR' IS NOT A VALID FORMAT FOR FILE 'REGISTRY'
'ABS' IS NOT A VALID FORMAT FOR FILE 'REGISTRY'

The following are valid formats:

Substance information can be displayed by requesting individual fields or predefined formats. The predefined substance formats are: (RN = CAS Registry Number)

REG - RN

SAM - Index Name, MF, and structure - no RNFIDE - All substance data, except sequence data

IDE - FIDE, but only 50 names
SQIDE - IDE, plus sequence data

SQIDE3 - Same as SQIDE, but 3-letter amino acid codes are used

SQD - Protein sequence data, includes RN

SQD3 - Same as SQD, but 3-letter amino acid codes are used

SQN - Protein sequence name information, includes RN

CALC - Table of calculated properties
EPROP - Table of experimental properties
PROP - EPROP and CALC

Any CA File format may be combined with any substance format to obtain CA references citing the substance. The substance formats must be cited first. The CA File predefined formats are:

ABS -- Abstract

APPS -- Application and Priority Information

BIB -- CA Accession Number, plus Bibliographic Data

CAN -- CA Accession Number

CBIB -- CA Accession Number, plus Bibliographic Data (compressed)

IND -- Index Data

IPC -- International Patent Classification

PATS -- PI, SO

STD -- BIB, IPC, and NCL

IABS --ABS, indented, with text labels

IBIB -- BIB, indented, with text labels

ISTD -- STD format, indented

OBIB ----- AN, plus Bibliographic Data (original) OIBIB ----- OBIB, indented with text labels

SBIB ----- BIB, no citations SIBIB ----- IBIB, no citations

The ALL format gives FIDE BIB ABS IND RE, plus sequence data when it is available.

The MAX format is the same as ALL.

The IALL format is the same as ALL with BIB ABS and IND indented, with text labels.

For additional information, please consult the following help messages:

HELP DFIELDS -- To see a complete list of individual display fields.

HELP FORMATS -- To see detailed descriptions of the predefined formats. ENTER DISPLAY FORMAT (IDE):ide

L3 ANSWER 1 OF 2 REGISTRY COPYRIGHT 2004 ACS on STN

RN 282095-40-3 REGISTRY

CN 1,2,3-Benzotriazine-3(4H)-butanoic acid, $4-oxo-\alpha-[2-oxo-2-[4'-(trifluoromethoxy)[1,1'-biphenyl]-4-yl]ethyl]- (9CI) (CA INDEX NAME)$

FS 3D CONCORD

MF C26 H20 F3 N3 O5

SR CA

LC STN Files: CA, CAPLUS

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 2 REFERENCES IN FILE CA (1907 TO DATE)
- 2 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- L3 ANSWER 2 OF 2 REGISTRY COPYRIGHT 2004 ACS on STN
- RN 282095-22-1 REGISTRY
- CN 2H-Isoindole-2-butanoic acid, α -[2-[4'-(acetyloxy)[1,1'-biphenyl]-4-yl]-2-oxoethyl]-1,3-dihydro-1,3-dioxo- (9CI) (CA INDEX NAME)
- FS 3D CONCORD
- MF C28 H23 N O7
- SR CA
- LC STN Files: CA, CAPLUS

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 2 REFERENCES IN FILE CA (1907 TO DATE)
- 2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

=> file caold

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION

274.09

0.44

FULL ESTIMATED COST

Patel

<3/12/2004>

FILE 'CAOLD' ENTERED AT 10:41:42 ON 12 MAR 2004
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FILE COVERS 1907-1966 FILE LAST UPDATED: 01 May 1997 (19970501/UP)

This file contains CAS Registry Numbers for easy and accurate substance identification. Title keywords, authors, patent assignees, and patent information, e.g., patent numbers, are now searchable from 1907-1966. TIFF images of CA abstracts printed between 1907-1966 are available in the PAGE display formats.

This file supports REG1stRY for direct browsing and searching of all substance data from the REGISTRY file. Enter HELP FIRST for more information.

=> s ll sss full

REG1stRY INITIATED

Substance data SEARCH and crossover from CAS REGISTRY in progress...
Use DISPLAY HITSTR (or FHITSTR) to directly view retrieved structures.

FULL SEARCH INITIATED 10:41:52 FILE 'REGISTRY' FULL SCREEN SEARCH COMPLETED - 343745 TO ITERATE

100.0% PROCESSED 343745 ITERATIONS SEARCH TIME: 00.00.13

2 ANSWERS

L5

2 SEA SSS FUL L1

L6 0 L5

=> file caplus
COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.42 430.35

FULL ESTIMATED COST

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<3/12/2004>

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FILE COVERS 1907 - 12 Mar 2004 VOL 140 ISS 12 FILE LAST UPDATED: 11 Mar 2004 (20040311/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

```
=> s 13
L7
             2 L3
=> s 14
\Gamma8
           126 L4
=> d 17 fbib hitstr abs total
T.7
     ANSWER 1 OF 2 CAPLUS COPYRIGHT 2004 ACS on STN
AN
     2001:247168 CAPLUS
DN
     134:266035
    Use of substituted 4-biarylbutyric and 5-biarylpentanoic acid derivatives
     for the treatment of multiple sclerosis
IN
     Fahrig, Thomas; Haning, Helmut; Riedl, Bernd; Braeunlich, Gabriele;
     Henning, Rolf
     Bayer Aktiengesellschaft, Germany
PA
SO
     PCT Int. Appl., 116 pp.
     CODEN: PIXXD2
DT
     Patent
LA
    English
FAN.CNT 1
     PATENT NO.
                      KIND
                           DATE
                                           APPLICATION NO. DATE
PΙ
    WO 2001022951
                      A2
                            20010405
                                           WO 2000-EP8890
                                                            20000912
    WO 2001022951
                      А3
                            20011011
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
             HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
             LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
             SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
             YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
             CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                           GB 1999-22710 A 19990924
    EP 1217994
                       A2
                            20020703
                                           EP 2000-965974 20000912
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, MC, IE, SI,
             LT, LV, FI, RO, MK, CY, AL
                                           GB 1999-22710 A 19990924
                                           WO 2000-EP8890 W 20000912
    JP 2003510272
                       T2
                            20030318
                                           JP 2001-526163 20000912
                                           GB 1999-22710 A 19990924
                                           WO 2000-EP8890 W 20000912
```

OS

IT

MARPAT 134:266035

282095-22-1P 282095-40-3P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of 4-biarylbutyric and 5-biarylpentanoic acid derivs. for the treatment of multiple sclerosis)

RN 282095-22-1 CAPLUS

CN 2H-Isoindole-2-butanoic acid, α -[2-[4'-(acetyloxy)[1,1'-biphenyl]-4-yl]-2-oxoethyl]-1,3-dihydro-1,3-dioxo-(9CI) (CA INDEX NAME)

RN 282095-40-3 CAPLUS

CN 1,2,3-Benzotriazine-3(4H)-butanoic acid, $4-\infty$ 0- α -[2- ∞ 0-2-[4'-(trifluoromethoxy)[1,1'-biphenyl]-4-yl]ethyl]- (9CI) (CA INDEX NAME)

AB The title compds. (T) xA-B-D-E-CO2H [I, A = aryl, heteroaryl; B = aryl, heteroaryl, bond; each T is a substituent group; x = 0, 1, or 2; D = CO, CH(OH); E = two or three carbon chain bearing one to three substituent groups which are independent or are involved in ring formation], useful for the treatment of multiple sclerosis, were prepared E.g., (rac)-2-[2-(1,3-dioxo-1,3-dihydro-2H-isoindol-2-yl)ethyl]-4-(4'-ethoxy[1,1'-biphenyl]-4-yl)-4-oxobutanoic acid was prepared Inhibitory activities of I against matrix metalloproteases was determined

L7 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2004 ACS on STN

AN 2000:475626 CAPLUS

DN 133:89429

TI Preparation of 4-aryl-4-oxo-2-(2-phthalimidoethyl)butanoates and analogs as matrix metalloprotease inhibitors

IN Fitzgerald, Mary F.; Gardiner, Philip J.; Nash, Kevin; Sturton, Graham; Benz, Gunter; Henning, Rolf; Schlemmer, Karl-Heinz; Riedl, Bernd; Haning, Helmut

PA Bayer Aktiengesellschaft, Germany

SO PCT Int. Appl., 146 pp.
CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO. KIND DATE APPLICATION NO. DATE

Patel

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PΙ
     WO 2000040539
                         A1
                              20000713
                                               WO 1999-EP10110 19991220
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              DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS,
              JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ,
              MD, RU, TJ, TM
          RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
              DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
              CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                               GB 1998-28845 A 19981230
                                               GB 1999-22709 A 19990924
     CA 2356053
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              IE, SI, LT, LV, FI, RO
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                                               GB 1999-22709 A 19990924
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     JP 2002534404
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                              20021015
                                               JP 2000-592250
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                                               GB 1998-28845 A 19981230
                                               GB 1999-22709 A 19990924
                                               WO 1999-EP10110W 19991220
     ZA 2001004651
                         Α
                              20020607
                                               ZA 2001-4651
                                                                 20010607
                                               GB 1998-28845 A 19981230
OS
     MARPAT 133:89429
TΤ
     282095-22-1P 282095-40-3P
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (preparation of 4-aryl-4-oxo-2-(2-phthalimidoethyl) butanoates and analogs as
        matrix metalloprotease inhibitors)
     282095-22-1 CAPLUS
```

RN 282095-22-1 CAPLUS CN 2H-Isoindole-2-butanoic acid, α -[2-[4'-(acetyloxy)[1,1'-biphenyl]-4-yl]-2-oxoethyl]-1,3-dihydro-1,3-dioxo- (9CI) (CA INDEX NAME)

RN 282095-40-3 CAPLUS
CN 1,2,3-Benzotriazine-3(4H)-butanoic acid, 4-oxo-α-[2-oxo-2-[4'-(trifluoromethoxy)[1,1'-biphenyl]-4-yl]ethyl]- (9CI) (CA INDEX NAME)

09869668

GΙ

AB RZZ1Z2CO2H [I; R = (un)substituted Ph or -heteroaryl; Z = bond, (un)substituted 1,4-phenylene, -heteroarylene; Z1 = CO, CH(OH), C(:NOH), etc.; Z2 = substituted (CH2)2-3] were prepared Thus, di-tert-Bu 2-(2-phthalimidoethyl)malonate was condensed with 4-(EtO)C6H4C6H4(COCH2Br)-4 (preparation each given) and the saponified product mono-decarboxylated to give

title compound II. Data for biol. activity of I were given.

RE.CNT 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> d his

(FILE 'HOME' ENTERED AT 10:33:14 ON 12 MAR 2004)

FILE 'REGISTRY' ENTERED AT 10:33:47 ON 12 MAR 2004

L1 STRUCTURE UPLOADED

L2 0 S L1

L3 2 S L1 SSS FULL

FILE 'MARPAT' ENTERED AT 10:34:49 ON 12 MAR 2004 L4 126 S L1 SSS FULL

FILE 'CAPLUS' ENTERED AT 10:40:39 ON 12 MAR 2004

FILE 'REGISTRY' ENTERED AT 10:41:17 ON 12 MAR 2004

FILE 'CAPLUS' ENTERED AT 10:41:24 ON 12 MAR 2004

FILE 'CAOLD' ENTERED AT 10:41:42 ON 12 MAR 2004 S L1

FILE 'REGISTRY' ENTERED AT 10:41:51 ON 12 MAR 2004 L52 S L1 SSS FULL FILE 'CAOLD' ENTERED AT 10:42:04 ON 12 MAR 2004 0 S L5 SSS FULL L6FILE 'CAPLUS' ENTERED AT 10:42:11 ON 12 MAR 2004 L7 2 S L3 $\Gamma8$ 126 S L4 => s respiratory disease and 1,1'-buphenyl MISMATCHED QUOTE '1,1'-BUPHENYL' Quotation marks (or apostrophes) must be used in pairs, one before and one after the expression you are setting off or masking. => s respiratory disease and phenyl 48 RESPIRATORY DISEASE AND PHENYL => s respiratory disease and 1,1'-biphenyl MISMATCHED QUOTE '1,1'-BIPHENYL' Quotation marks (or apostrophes) must be used in pairs, one before and one after the expression you are setting off or masking. => s respiratory disease and biphenyl 8 RESPIRATORY DISEASE AND BIPHENYL => s 17 and 18 2 L7 AND L8 L11=> s 18 and respiratory diseases 0 L8 AND RESPIRATORY DISEASES => s 19 and 110 L134 L9 AND L10 => s 18 and 113 0 L8 AND L13 => s 18 and 110 0 L8 AND L10 L15=> d l13 fbib hitstr abs total L13 ANSWER 1 OF 4 CAPLUS COPYRIGHT 2004 ACS on STN 2003:242154 CAPLUS ΑN 138:254964 DN TIPreparation of 4-(2-amino-1-hydroxyethyl)-2-(hydroxymethyl)phenols as selective $\beta 2$ -adrenoreceptor agonists for treatment of respiratory diseases Box, Philip Charles; Coe, Diane Mary; Looker, Brian Edgar; Procopiou, INPanayiotis Alexandrou PΑ Glaxo Group Limited, UK SO PCT Int. Appl., 117 pp. CODEN: PIXXD2 DTPatent

GB 2001-26997 A 20011109

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LA English FAN.CNT 1
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I AIN .	PATENT NO.				KIND DATE					APPLICATION NO.					DATE			
ΡI	WO	0 2003024439			A1 20030327				W	20	02-G	B414	0	20020911				
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			CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,
			GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,	LK,	LR,
			LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	OM,	PH,
			PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	ТJ,	TM,	TN,	TR,	TT,	TZ,
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			RU,	ТJ,	TM													
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			PT,	SE,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,
			NE,	SN,	TD,	TG												
									GB 2001-22201 A 2001091									

OS MARPAT 138:254964

HO

$$N \longrightarrow 0$$
 $N \longrightarrow 0$
 $N \longrightarrow N$
 $N \longrightarrow N$

AB Title phenethanolamines I [wherein m = 2-8, n = 2-5; and m + n = 4-10; R1 = H, (halo)alkyl, OH, halo, XCONR9R10, XNR8COR9, XNR8CONR9R10, XNR8SO2R9, XSO2NR11R12, XNR8SO2NR9R10, XNR9R10, XN+R8R9R10, XNR8CO2R9, XCO2R9, XNR8CONR8CONR9R10, XSR9, XSOR9, XSO2R9, or (un)substituted X-(hetero)aryl, or X-aryloxy; R2 and R3 = independently H, OH, (halo)alkyl, (halo)alkoxy, halo, or aryl(alkyl); R4 and R5 = independently H or alkyl with the proviso that the total number of C atoms in R4 and R5 ≤ 4; R6 and R7 = independently H or alkyl with the proviso that the total number of C atoms in R6 and R7 ≤ 4; X = (CH2)p or alkenylene; p = 0-6; R8 and R9 = independently H, or (un)substituted (cyclo)alkyl, (hetero)aryl, or (hetero)arylalkyl; R10 = H or (cyclo)alkyl; R11 and R12 = independently H, (cyclo)alkyl, (hetero)aryl, or (hetero)arylalkyl; or NR11R12 = (un)substituted heterocyclyl; with addnl. provisos; or salts, solvates, or

physiol. functional derivs. thereof] were prepared as selective stimulants of β 2-adrenoceptors. For example, coupling of (5R)-5-(2,2-dimethyl-4H-1, 3-benzodioxin-6-y1) -1, 3-oxazolidin-2-one with [2-[(6bromohexyl)oxy]ethoxy](tert-butyl)dimethylsilane using NaH in DMF (preparation of starting materials given), deprotection of the alc. with Bu4NF in THF, etherification with 3-nitrobenzyl bromide in DMF in the presence of NaH, and hydrogenation over platinum oxide afforded (5R)-3-[6-[2-[(3aminobenzyl)oxy]ethoxy]hexyl]-5-(2,2-dimethyl-4H-1,3-benzodioxin-6-yl)-1,3oxazolidin-2-one. Reaction of the amine with PhNCO in i-PrOH to give the urea, ring opening using Me3SiOK in THF, and hydrolysis with AcOH and H2O provided (R)-II \bullet AcOH. The latter exhibited β 2-adrenoreceptor agonist activity with IC50 values $< 1 \mu M$ and showed five fold selectivity for the $\beta2$ - over $\beta3$ -adrenoceptors. Elec.- or spasmogen-induced contractions in human or guinea pig airway tissue were typically suppressed by preferred compds. of the invention in < 30 min and maintained for > 3 h. Thus, I are useful for the prophylaxis and treatment of respiratory diseases.

RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

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L13 ANSWER 2 OF 4 CAPLUS COPYRIGHT 2004 ACS on STN
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AN 2002:658074 CAPLUS

DN 137:201142

TI Preparation of 2-hydroxy-2-[4-hydroxy-3-(hydroxymethyl)**phenyl**]ethanamine derivatives as β2-adrenoreceptor agonists for treatment of **respiratory diseases**

IN Biggadike, Keith; Coe, Diane Mary; Edney, Dean David; Halton, Abigail; Looker, Brian Edgar; Monteith, Michael John; Moore, Rebecca Jane; Patel, Rajnikant; Procopiou, Panayiotis Alexandrou

PA Glaxo Group Limited, UK

SO PCT Int. Appl., 88 pp. CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

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PATENT NO.
                      KIND
                            DATE
                                           APPLICATION NO.
                                                             DATE
PI
     WO 2002066422
                       A1
                            20020829
                                           WO 2002-EP1387
                                                             20020211
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                                           GB 2001-3630
                                                          A 20010214
                                           GB 2001-26998 A 20011109
     EP 1360174
                       A1
                            20031112
                                           EP 2002-706735
                                                            20020211
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                                           GB 2001-3630
                                                          A 20010214
                                           GB 2001-26998 A 20011109
                                           WO 2002-EP1387 W 20020211
    NO 2003003594
                                           NO 2003-3594
                       Α
                            20031002
                                                            20030813
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GB 2001-3630 A 20010214 GB 2001-26998 A 20011109 WO 2002-EP1387 W 20020211

OS MARPAT 137:201142

GΙ

HO—CH₂
OH
$$R^4$$
 R^5
 R^3
 R^3
 R^3

HO
$$-CH_2$$
OH
 $-N$
 $-SO_2=NH_2-$
II

AΒ The title phenethanolamines I [wherein m = 2-8; n = 3-11; with the proviso the m + n = 5-19; R1 = XSO2NR6R7; R2 and R3 = independently H,(halo)alkyl, alkoxy, halo, or Ph; R4 and R5 = independently H or alkyl with the proviso that the total number of C's in R4 and R5 \leq 4; X = (CH2)p or alkenylene; p = 0-6; R6 and R7 = independently H, CONR8R9, or (un) substituted (cyclo) alkyl, Ph, or phenylalkyl; or NR6R7 = (un) substituted N-containing ring; R8 and R9 = independently H, (cyclo) alkyl, Ph, or phenylalkyl; or salts, solvates, or physiol. functional derivs. thereof] were prepared as β 2-adrenoreceptor agonists. For example, (R)-II•AcOH was synthesized in ten steps beginning with the coupling of di-t-Bu iminodicarboxlyate with 2-bromo-1-(2,2-dimethyl-4H-1,3benzodioxin-6-yl)ethanone in the presence of Cs2CO3 in AcCN. Forty-nine compds. of the invention displayed agonist activity against human $\beta2\text{-adrenoreceptors}$ with IC50 values below 1 $\mu M.$ Thus, I and pharmaceutical compns. containing them are useful in the prophylaxis and treatment of respiratory diseases (no data).

RE.CNT 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L13 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2004 ACS on STN

AN 2002:391690 CAPLUS

DN 136:386115

TI Substituted 2-phenylaminoimidazoline **phenyl** ketone derivatives as human platelet IP receptor antagonists

IN Jahangir, Alam

PA F. Hoffmann-La Roche Ag, Switz.

SO PCT Int. Appl., 59 pp. CODEN: PIXXD2

Patel

<3/12/2004>

Patent

DT

LA English FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. DATE _____ WO 2001-EP12776 20011105 PIWO 2002040453 **A**1 20020523 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG US 2000-248888PP 20001114 AU 2002021808 Α5 20020527 AU 2002-21808 US 2000-248888PP 20001114 WO 2001-EP12776W 20011105 BR 2001015291 20030819 BR 2001-15291 20011105 US 2000-248888PP 20001114 WO 2001-EP12776W 20011105 20030903 EP 1339694 **A**1 EP 2001-996531 20011105 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR US 2000-248888PP 20001114 WO 2001-EP12776W 20011105 US 6417186 В1 20020709 US 2001-14110 20011113 US 2002091147 20020711 **A**1

20030513

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NO 2003002142

MARPAT 136:386115

OS

GΙ

US 2000-248888PP 20001114

US 2000-248888PP 20001114 WO 2001-EP12776W 20011105

20030513

NO 2003-2142

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AΒ
     Title compds. I [] were prepared For instance, 4-fluoroacetophenone and
     4-nitrobenzaldehyde were reacted together (EtOHaq, KOH) to give
     1-[4-fluorophenyl]-3-[4-nitrophenyl]propenone. This intermediate was
     reduced (EtOAc, H2-10% Pd/C) and reacted with 2-chloro-4,5-dihydro-1H-
     imidazole sulfate to give II in 54.2\% overall yield. Example compds. had
     pKi in the range of 7.1 to 9.6 for the human platelet IP receptor; II had
     pKi = 9.50. I are used for the treatment of diseases associated with pain,
     inflammation, urinary tract disease states, respiratory
     disease states, edema formation, or hypotensive vascular diseases.
RE.CNT 3
              THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
    ANSWER 4 OF 4 CAPLUS COPYRIGHT 2004 ACS on STN
L13
AN
     2001:693269 CAPLUS
DN
     135:257467
ΤI
     Preparation of N-(arylmethoxycarbonyl)phenylalanine derivatives as IP
     antagonists
IN
     Cournoyer, Richard Leo; Keitz, Paul Francis; Lowrie, Lee Edwin, Jr.;
     Muehldorf, Alexander Victor; O'Yang, Counde; Yasuda, Dennis Mitsugu
PA
     F. Hoffmann-La Roche A.-G., Switz.
SO
     PCT Int. Appl., 100 pp.
     CODEN: PIXXD2
^{-}D\mathbf{T}^{-}
    Patent
LА
     English
FAN.CNT 1
     PATENT NO.
                      KIND DATE
                                           APPLICATION NO. DATE
PΙ
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                     A1
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                                           US 2000-190129PP 20000316
                                           US 2000-247129PP 20001110
     BR 2001009235
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     EP 1265853
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                                           EP 2001-925395
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                                           WO 2001-EP2597 W 20010308
    US 2001056100
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                            20011227
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                                                            20010314
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Patel <3/12/2004>

20040217

US 2000-190129PP 20000316

B2

US 6693098

09869668

US 2000-247129PP 20001110 NO 2002004387 20021021 NO 2002-4387 20020913 US 2000-190129PP 20000316 US 2000-247129PP 20001110 WO 2001-EP2597 W 20010308 US 2003220367 A1 20031127 US 2003-434809 20030509 US 2000-190129PP 20000316 US 2000-247129PP 20001110 US 2001-810436 A320010314 OS MARPAT 135:257467

Page 19

Ι

OS MARPAT 135:257467

$$_{R1}^{A}$$
 $_{R2}^{A}$ $_{R2}^{(CH_2)}$ $_{M}^{m}$ $_{H}^{(CH_2)}$ $_{R4}^{n}$ $_{R3}^{(CH_2)}$

AΒ Title compds. I [wherein R1, R2, and R3 = independently (un) substituted (hetero)aryl; R4 = COOH or tetrazolyl; A = single bond, O(CH2)q, S(CH2)q, NR'(CH2)q, (CH2)qO, O(CH2)qO, (CH2)qO(CH2)q, (CH2)nCO(CH2)q, CONH, (CH2)q, CH:CH, or C.tplbond.C; R' = H or alkyl; B = (CH2)r, CH2O, CH2OCH2, or CH2NH; m, q, and r = independently 1-3; n and p = independently 0-3; or individual isomers, racemic or nonracemic mixts. of isomers, or pharmaceutically acceptable salts or solvates thereof] were prepared as prostaglandin IP receptor antagonists. For example, 4-vinylbenzoic acid was esterified with MeOH (97.4%). The ester was hydroborated with 9-BBN in THF and oxidized with alkaline HOOH to give 4-(2-hydroxyethyl)benzoic acid Me ester (64.7%). Etherification with PhOH in the presence of PPh3 and diEt azodicarboxylate in THF (16.8%), followed by reduction with LiAlH4, condensation with Me (R)-2-isocyanato-3-phenylpropionate, and hydrolysis, afforded II. The latter showed affinity toward the human platelet IP receptor with a Ki value of 6.6. I are useful for the treatment of inflammatory conditions, pain, bladder disorders, hypotensive vascular diseases, and respiratory diseases, such as allergies and asthma (no data).

RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

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FILE 'REGISTRY' ENTERED AT 10:33:47 ON 12 MAR 2004

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FILE 'MARPAT' ENTERED AT 10:34:49 ON 12 MAR 2004

L4 126 S L1 SSS FULL

FILE 'CAPLUS' ENTERED AT 10:40:39 ON 12 MAR 2004

FILE 'REGISTRY' ENTERED AT 10:41:17 ON 12 MAR 2004

FILE 'CAPLUS' ENTERED AT 10:41:24 ON 12 MAR 2004

FILE 'CAOLD' ENTERED AT 10:41:42 ON 12 MAR 2004 S L1

FILE 'REGISTRY' ENTERED AT 10:41:51 ON 12 MAR 2004 L5 2 S L1 SSS FULL

FILE 'CAOLD' ENTERED AT 10:42:04 ON 12 MAR 2004 L6 0 S L5 SSS FULL

FILE 'CAPLUS' ENTERED AT 10:42:11 ON 12 MAR 2004

L7 2 S L3

L8 126 S L4

L9 48 S RESPIRATORY DISEASE AND PHENYL

L10 8 S RESPIRATORY DISEASE AND BIPHENYL

L11 2 S L7 AND L8

L12 0 S L8 AND RESPIRATORY DISEASES

L13 4 S L9 AND L10

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L15 0 S L8 AND L10

=> s 18 and indilinone

L16 0 L8 AND INDILINONE

=> s 18 and inmidazolidinone

L17 0 L8 AND INMIDAZOLIDINONE

=> s 18 and imidazolidinone

L18 0 L8 AND IMIDAZOLIDINONE

=> s 18 and phtahlazine

L19 0 L8 AND PHTAHLAZINE

=> s 18 and asthma

L20 8 L8 AND ASTHMA

=> s 18 and ARDS

L21 0 L8 AND ARDS

=> a 18 and TNF alpha

Patel

09869668

Page 21 A IS NOT A RECOGNIZED COMMAND The previous command name entered was not recognized by the system. For a list of commands available to you in the current file, enter "HELP COMMANDS" at an arrow prompt (=>). => s 18 and TNF alpha L22 2 L8 AND TNF ALPHA => s 16 and rhinitis 0 L6 AND RHINITIS => s 18 and bronchiectasis 0 L8 AND BRONCHIECTASIS => s 18 and silicosis 0 L8 AND SILICOSIS => s silicosis and bronchiectasis 15 SILICOSIS AND BRONCHIECTASIS => s 116 and 18 0 L16 AND L8 => s 126 and asthma 11 L26 AND ASTHMA L28 => d his (FILE 'HOME' ENTERED AT 10:33:14 ON 12 MAR 2004) FILE 'REGISTRY' ENTERED AT 10:33:47 ON 12 MAR 2004 L1STRUCTURE UPLOADED L2 0 S L1 L32 S L1 SSS FULL FILE 'MARPAT' ENTERED AT 10:34:49 ON 12 MAR 2004 L4126 S L1 SSS FULL FILE 'CAPLUS' ENTERED AT 10:40:39 ON 12 MAR 2004 FILE 'REGISTRY' ENTERED AT 10:41:17 ON 12 MAR 2004 FILE 'CAPLUS' ENTERED AT 10:41:24 ON 12 MAR 2004 FILE 'CAOLD' ENTERED AT 10:41:42 ON 12 MAR 2004 S L1 FILE 'REGISTRY' ENTERED AT 10:41:51 ON 12 MAR 2004 L52 S L1 SSS FULL

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FILE 'CAPLUS' ENTERED AT 10:42:11 ON 12 MAR 2004

L72 S L3 126 S L4 $\Gamma8$

1.9 48 S RESPIRATORY DISEASE AND PHENYL

Patel